PTO-1390 (Rev. 07-2005) Approved for use through 03/31/2007. OMB 0651-0021 U. S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. ATTORNEY'S DOCKET NUMBER TRANSMITTAL LETTER TO THE UNITED STATES 0113847.00127US1 DESIGNATED/ELECTED OFFICE (DO/EO/US) 10 10 (i/kn 201, see 37 CFR 1.5) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371 INTERNATIONAL APPLICATION NO. INTERNATIONAL FILING DATE RIORITY DATE CLAIMED PCT/EP2005/002086 28 February 2005 26 February 2004 TITLE OF INVENTION NOVEL STEROID AGONIST FOR FXR APPLICANT(S) FOR DO/EO/US Roberto PELLICCIARI Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information: This is a FIRST submission of items concerning a submission under 35 U.S.C. 371. This is a SECOND or SUBSEQUENT submission of items concerning a submission under 35 U.S.C. 371. This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below. The US has been elected (Article 31). 4. x A copy of the International Application as filed (35 U.S.C. 371 (c)(2)) 5. x is attached hereto (required only if not communicated by the International Bureau). has been communicated by the International Bureau. is not required, as the application was filed in the United States Receiving Office (RO/US). An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)). is attached hereto. has been previously submitted under 35 U.S.C. 154(d)(4). Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3)) are attached hereto (required only if not communicated by the International Bureau). have been communicated by the International Bureau. have not been made; however, the time limit for making such amendments has NOT expired. have not been made and will not be made. An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)). An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)). An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)). Items 11 to 20 below concern document(s) or information included: An Information Disclosure Statement under 37 CFR 1.97 and 1.98. 12 An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included. 13 A preliminary amendment. 14. х An Application Data Sheet under 37 CFR 1.76. 15 A substitute specification. 16 A power of attorney and/or change of address letter. 17 A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 – 1.825. 18 A second copy of the published International Application under 35 U.S.C. 154(d)(4). 19 A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).

IAP9 Rec'd PCT/PTO 25 AUG 2006

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U.S.ADIC/TE	390°8°4	37 G R 1.5)	INTERNATIONAL APPLICATION NO. PCT/EP2005/002086		ATTORNEY'S DOCKET NUMBER 0113847.00127US1			
20. x Other items or information: PCT Request PCT Publication WO 2005/082925 A2 PCT Publication WO 2005/082925 A3 (with Search Report)								
The following fees have been submitted						CALCULATION	S PTO USEONLY	
21. x Basic national fee (37 CFR 1.492(a))						\$ 300.0		
22. x Examination fee (37 CFR 1.492(c)) If the written opinion prepared by ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4)						\$ 200.0	0	
23. X Search fee (37 CFR 1.492(b)) If the written opinion of the ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4)						\$ 400.0	0	
TOTAL OF 21, 22 and 23 =						\$ 900.0	0	
Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing in compliance with 37 CFR 1.821(c) or (e) or computer program listing in an electronic medium) (37 CFR 1.492(j)). The fee is \$250 for each additional 50 sheets of paper or fraction thereof.								
Total Sheets	Extra Sheets		additional 50 or fraction up to a whole number)	1	RATE			
20 - 100 =	/50 =		x \$250.00		x \$250.00	\$		
Surcharge of \$130 for furnishing any of the search fee, examination fee, or the oath or declaration after the date of commencement of the national stage (37 CFR 1.492(h)).						\$		
CLAIMS NUMBER FILED		MBER FILED	NUMBER EXTRA RATE					
Total claims 30 - 20		0 - 20 =	10 x 25.00		500.0	0		
Independent claims 3 - 3 =			x		0.0			
MULTIPLE DEP	ENDENT CLAIM(S) (if applicable)		+	180.00	360.00		
TOTAL OF ABOVE CALCULATIONS =						\$ 1,760.0	0	
X Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by ½.								
SUBTOTAL =						\$ 880.00		
Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(i)).						\$		
TOTAL NATIONAL FEE =						\$ 880.00		
Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property +						\$		
						\$		
TOTAL FEES ENCLOSED =					\$ 880.00			
	· · · · · · · · · · · · · · · · · · ·	<u></u>				Amount to be refunded:	\$	
						Amount to be charged	\$	
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a. A check in the amount of \$ to cover the above fees is enclosed.								
b. x Please charge my Deposit Account No. 08-0219 in the amount of \$ 880.00 to cover the above fees. A duplicate copy of this sheet is enclosed.								
c. x The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 08-0219 . A duplicate copy of this sheet is enclosed.								
d. Fees are to be charged to a credit card. WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038								
NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed an granted to restore the international Application to pending status.								
med an granted to restore the international Application to ponding status.								
SEND ALL CORRESPONDENCE TO: SIGNATURE SIGNATURE								
Jamie T. Wisz								
CUSTOMER NUMBER: 24395								
58,429								
REGISTRATION NUMBER								
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10/590848 IAP9 Rec'd PCT/PTO 25 AUG 2006

PTO/SB/17 (01-06)
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Under the Paperwork Reduction Act of 1995, no person are required to respond to a collection of information unless it displays a valid OMB control number. Complete if Known Fees pursuant to the Consolidated Appropriations Act, 2005 (H.R. 4818). **Application Number** Not Yet Assigned FEE TRANSMITTAL Filing Date For FY 2006 Roberto PELLICCIARI First Named Inventor Not Yet Assigned **Examiner Name** Applicant claims small entity status. See 37 CFR 1.27 Art Unit 0113847.00127US1 TOTAL AMOUNT OF PAYMENT 880.00 Attorney Docket No. METHOD OF PAYMENT (check all that apply) Money Order None Other (please identify): Check Credit Card Wilmer Cutler Pickering Hale and Dorr LLP x Deposit Account Deposit Account Number: 08-0219 Deposit Account Name: For the above-identified deposit account, the Director is hereby authorized to: (check all that apply) Charge fee(s) indicated below, except for the filing fee x Charge fee(s) indicated below Charge any additional fee(s) or underpayment of Credit any overpayments fee(s) under 37 CFR 1.16 and 1.17 FEE CALCULATION (All the fees below are due upon filing or may be subject to a surcharge.) 1. BASIC FILING, SEARCH, AND EXAMINATION FEES **FILING FEES SEARCH FEES EXAMINATION FEES Small Entity Small Entity** Small Entity Fees Paid (\$) Fee (\$) Fee (\$) **Application Type** Fee (\$) Fee (\$) Fee (\$) Fee (\$) 200 100 300 150 500 250 Utility 200 100 100 50 130 65 Design 300 160 80 Plant 200 100 150 Reissue 300 150 500 250 600 300 0 0 0 Provisional 200 100 0 Small Entity 2. EXCESS CLAIM FEES Fee (\$) Fee (\$) Fee Description 25 Each claim over 20 (including Reissues) 50 200 100 Each independent claim over 3 (including Reissues) 180 Multiple dependent claims 360 **Total Claims** Extra Claims Fee Paid (\$) **Multiple Dependent Claims** Fee (\$) - 20 = 10 Fee (\$) Fee Paid (\$) HP = highest numer of total claims paid for, if greater than 20. Indep. Claims Extra Claims Fee Paid (\$) Fee (\$) 3 - 3 = HP = highest numer of independent claims paid for, if greater than 3. If the specification and drawings exceed 100 sheets of paper (excluding electronically filed sequence or computer listings under 37 CFR 1.52(e)), the application size fee due is \$250 (\$125 for small entity) for each additional 50 sheets or fraction thereof. See 35 U.S.C. 41(a)(1)(G) and 37 CFR 1.16(s). Fee Paid (\$) Extra Sheets Number of each additional 50 or fraction thereof Fee (\$) 20 - 100 = (round up to a whole number) x 4. OTHER FEE(S) Fees Paid (\$) Non-English Specification, \$130 fee (no small entity discount) Other (e.g., late filing surcharge): 1631 Basic National Stage fee 150.00 1633 National Stage Examination Fee - all other ... 100.00 1632 National Stage Search Fee - all other situations 200.00 1615 National Stage claims - extra total (over twenty) 250.00 180.00 1616 National Stage claims - multiple dependent SUBMITTED BY Registration No. Wis mee 58.429 Telephone (202) 663-6000 Signature August 25, 2006 Name (Print/Type) Jamie T. Wisz